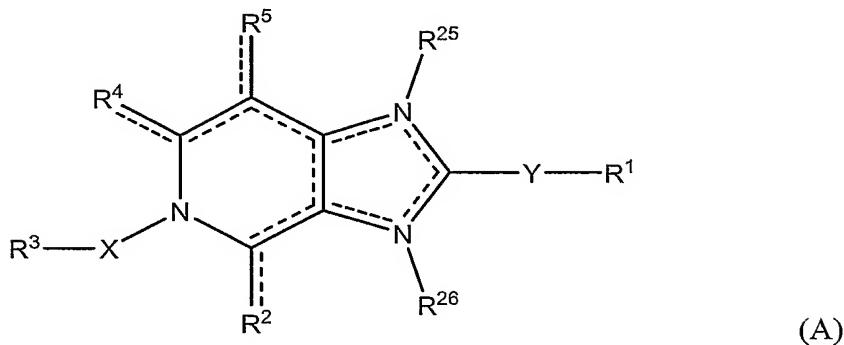


5

We claim:

1. A compound having the general formula (A),



10 wherein:

the dotted lines represent an optional double bond, provided that no two double bonds are adjacent to one another, and that the dotted lines represent at least 3, optionally 4 double bonds;

15 R^1 is selected from hydrogen, aryl, heterocyclic, $C_{1-C_{10}}$ alkoxy, $C_{1-C_{10}}$ thioalkyl, $C_{1-C_{10}}$ alkyl-amino, $C_{1-C_{10}}$ dialkyl-amino, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, and C_{4-10} cycloalkynyl, wherein each are optionally substituted with 1 or more R^6 ;

20 Y is selected from single bond, O, $S(O)_m$, NR^{11} , or C_{1-10} alkylene, C_{2-10} alkenylene, C_{2-10} alkynylene, wherein each may optionally include 1 to 3 heteroatoms selected from O, S or N;

25 R^2 and R^4 are independently selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, halogen, -OH, -CN, $-NO_2$, $-NR^7R^8$, haloalkyloxy, haloalkyl, $-C(=O)R^9$, $-C(=S)R^9$, SH, aryl, aryloxy, arylthio, arylalkyl, C_{1-18} hydroxyalkyl, C_{3-10} cycloalkyl, C_{3-10} cycloalkyloxy, C_{3-10} cycloalkylthio, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, or heterocyclic, provided that when one of R^{25} or R^{26} is present, then either R^2 or R^4 is selected from $(=O)$, $(=S)$, and $=NR^{27}$;

30 X is selected from $C_{1-C_{10}}$ alkylene, C_{2-10} alkenylene or C_{2-10} alkynylene, where each may include one or more heteroatoms selected from O, S, or N, provided any such heteroatom is not adjacent to the N in the ring;

m is any integer from 0 to 2;

5 R³ is selected from aryl, aryloxy, arylthio, cycloalkyl, cycloalkenyl, cycloalkynyl, aryl-N(R¹⁰)-, or heterocyclic, where each said substituent may be optionally substituted with at least one R¹⁷, provided that for cycloalkenyl the double bond is not adjacent to a nitrogen, and provided R³ M-Q- is not biphenyl;

10 R⁵ is selected from hydrogen; C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, C₁₋₁₈ alkoxy, C₁₋₁₈ alkylthio, halogen, -OH, -CN, -NO₂, -NR⁷R⁸, haloalkyloxy, haloalkyl, -C(=O)R⁹, -C(=O)OR⁹, -C(=S)R⁹, SH, aryl, aryloxy, arylthio, arylalkyl, C₁₋₁₈ hydroxyalkyl, C₃₋₁₀ cycloalkyl, C₃₋₁₀ cycloalkyloxy, C₃₋₁₀ cycloalkylthio, C₃₋₁₀ cycloalkenyl, C₇₋₁₀ cycloalkynyl, or heterocyclic;

15 R⁶ is selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, C₁₋₁₈ alkoxy, C₁₋₁₈ alkylthio, C₁₋₁₈ alkylsulfoxide, C₁₋₁₈ alkylsulfone, C₁₋₁₈ halo-alkyl, C₂₋₁₈ halo-alkenyl, C₂₋₁₈ halo-alkynyl, C₁₋₁₈ halo-alkoxy, C₁₋₁₈ halo-alkylthio, C₃₋₁₀ cycloalkyl, C₃₋₁₀ cycloalkenyl, C₇₋₁₀ cycloalkynyl, halogen, OH, CN, cyanoalkyl, -CO₂R¹⁸, NO₂, -NR⁷R⁸, C₁₋₁₈ haloalkyl, C(=O)R¹⁸, C(=S)R¹⁸, SH, aryl, aryloxy, arylthio, arylsulfoxide, arylsulfone, arylsulfonamide, aryl(C₁₋₁₈)alkyl, aryl(C₁₋₁₈)alkyloxy, aryl(C₁₋₁₈)alkylthio, heterocyclic, C₁₋₁₈ hydroxyalkyl, where each may be optionally substituted with at least 1 R¹⁹;

20 R⁷ and R⁸ are independently selected from hydrogen, C₁₋₁₈ alkyl, C₁₋₁₈ alkenyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, heterocyclic, -C(=O)R¹²; -C(=S)R¹², an amino acid residue linked through a carboxyl group thereof, or where R⁷ and R⁸ together with the nitrogen form a heterocyclic;

25 R⁹ and R¹⁸ are independently selected from hydrogen, OH, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, C₁₋₁₈ alkoxy, -NR¹⁵R¹⁶, aryl, an amino acid residue linked through an amino group of the amino acid, CH₂OCH(=O)R^{9a}, or CH₂OC(=O)OR^{9a} where R^{9a} is C_{1-C₁₂} alkyl, C_{6-C₂₀} aryl, C_{6-C₂₀} alkylaryl or C_{6-C₂₀} aralkyl;

30 R¹⁰ and R¹¹ are independently selected from the group consisting of hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, aryl, -C(=O)R¹², heterocyclic, or an amino acid residue;

35 R¹² is selected from the group consisting of hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, or an amino acid residue;

5 R¹⁵ and R¹⁶ are independently selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, or an amino acid residue;

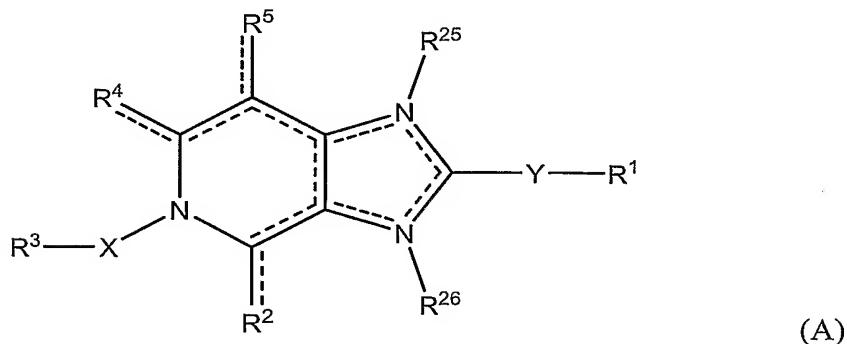
10 R¹⁷ is independently M-Q- wherein M is a ring optionally substituted with 1 or more R¹⁹, and Q is a bond or a linking group connecting M to R³ having 1 to 10 atoms and optionally substituted with 1 or more R¹⁹;

15 R¹⁹ is selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, C₁₋₁₈ alkoxy, C₂₋₁₈ alkenyloxy, C₂₋₁₈ alkynyoxy, C₁₋₁₈ alkylthio, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, C₄₋₁₀ cycloalkynyl, halogen, -OH, -CN, cyanoalkyl, -NO₂, -NR²⁰R²¹, C₁₋₁₈ haloalkyl, C₁₋₁₈ haloalkyloxy, -C(=O)R¹⁸, -C(=O)OR¹⁸, -OalkenylC(=O)OR¹⁸, -OalkylC(=O)NR²⁰R²¹, -OalkylOC(=O)R¹⁸, -C(=S)R¹⁸, SH, -C(=O)N(C₁₋₆ alkyl), -N(H)S(O)(O)(C₁₋₆ alkyl), aryl, heterocyclic, C₁₋₁₈ alkylsulfone, arylsulfoxide, arylsulfonamide, aryl(C₁₋₁₈)alkyloxy, aryloxy, aryl(C₁₋₁₈ alkyl)oxy, arylthio, aryl(C₁₋₁₈)alkylthio or aryl(C₁₋₁₈)alkyl, where each may be optionally substituted with 1 or more =O, NR²⁰R²¹, CN, C₁₋₁₈ alkoxy, heterocyclic, C₁₋₁₈ haloalkyl, heterocyclic alkyl, 20 heterocyclic connected to R¹⁷ by alkyl, alkoxyalkoxy or halogen;

20 R²⁰ and R²¹ are independently selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, -C(=O)R¹², or -C(=S)R¹²;

25 R²⁷ is selected from hydrogen, C₁₋₁₈ alkyl, C₃₋₁₀ cycloalkyl, (C₃₋₁₀ cycloalkyl)-C₁₋₆ alkyl, aryl, and aryl C₁₋₁₈ alkyl, and salts, tautomers, isomers and solvates thereof.

2. A compound having the general formula (A),



wherein:

5 the dotted lines represent an optional double bond, provided that no two double bonds are adjacent to one another, and that the dotted lines represent at least 3, optionally 4 double bonds;

10 R¹ is selected from hydrogen, aryl, heterocyclic, C₁-C₁₀ alkoxy, C₁-C₁₀ thioalkyl, C₁-C₁₀ alkyl-amino, C₁-C₁₀ dialkyl-amino, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, and C₄₋₁₀ cycloalkynyl, wherein each are optionally substituted with 1 or more R⁶;

15 Y is selected from single bond, O, S(O)_m, NR¹¹, or C₁₋₁₀ alkylene, C₂₋₁₀ alkenylene, C₂₋₁₀ alkynylene, wherein each may optionally include 1 to 3 heteroatoms selected from O, S or N;

20 R² and R⁴ are independently selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, C₁₋₁₈ alkoxy, C₁₋₁₈ alkylthio, halogen, -OH, -CN, -NO₂, -NR⁷R⁸, haloalkyloxy, haloalkyl, -C(=O)R⁹, -C(=S)R⁹, SH, aryl, aryloxy, arylthio, arylalkyl, C₁₋₁₈ hydroxyalkyl, C₃₋₁₀ cycloalkyl, C₃₋₁₀ cycloalkyloxy, C₃₋₁₀ cycloalkylthio, C₃₋₁₀ cycloalkenyl, C₇₋₁₀ cycloalkynyl, or heterocyclic, provided that when one of R²⁵ or R²⁶ is present, then either R² or R⁴ is selected from (=O), (=S), and =NR²⁷;

25 X is selected from C₁-C₁₀ alkylene, C₂₋₁₀ alkenylene or C₂₋₁₀ alkynylene, where each may include one or more heteroatoms selected from O, S, or N, provided any such heteroatom is not adjacent to the N in the ring;

m is any integer from 0 to 2;

30 R³ is a heterocycle optionally substituted with at least one R¹⁷ provided, however, that R³ optionally substituted with at least one R¹⁷ is not pyridinyl or 5-chlorothienyl, provided that R³-MQ is not biphenyl;

R⁵ is selected from hydrogen; C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, C₁₋₁₈ alkoxy, C₁₋₁₈ alkylthio, halogen, -OH, -CN, -NO₂, -NR⁷R⁸, haloalkyloxy, haloalkyl, -C(=O)R⁹, -C(=O)OR⁹, -C(=S)R⁹, SH, aryl, aryloxy, arylthio, arylalkyl, C₁₋₁₈ hydroxyalkyl, C₃₋₁₀ cycloalkyl, C₃₋₁₀ cycloalkyloxy, C₃₋₁₀ cycloalkylthio, C₃₋₁₀ cycloalkenyl, C₇₋₁₀ cycloalkynyl, or heterocyclic;

35 R⁶ is selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, heterocyclic, C₁₋₁₈ alkoxy, C₁₋₁₈ alkylthio, C₁₋₁₈ alkylsulfoxide, C₁₋₁₈ alkylsulfone, C₁₋₁₈ halo-alkyl, C₂₋₁₈ halo-alkenyl, C₂₋₁₈ halo-alkynyl, C₁₋₁₈ halo-alkoxy, C₁₋₁₈ halo-alkylthio, C₃₋₁₀ cycloalkyl, C₃₋₁₀ cycloalkenyl, C₇₋₁₀ cycloalkynyl, halogen, OH, CN,

5 cyanoalkyl, $-\text{CO}_2\text{R}^{18}$, NO_2 , $-\text{NR}^7\text{R}^8$, C_{1-18} haloalkyl, $\text{C}(=\text{O})\text{R}^{18}$, $\text{C}(=\text{S})\text{R}^{18}$, SH , aryl, aryloxy, arylthio, arylsulfoxide, arylsulfone, arylsulfonamide, aryl(C_{1-18})alkyl, aryl(C_{1-18})alkyloxy, aryl(C_{1-18})alkylthio, C_{1-18} hydroxyalkyl, where each may be optionally substituted with at least 1 R^{19} ;

10 R^7 and R^8 are independently selected from hydrogen, C_{1-18} alkyl, C_{1-18} alkenyl, aryl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, heterocyclic, $-\text{C}(=\text{O})\text{R}^{12}$; $-\text{C}(=\text{S})\text{R}^{12}$, an amino acid residue linked through a carboxyl group thereof, or where R^7 and R^8 together with the nitrogen form a heterocyclic;

15 R^9 and R^{18} are independently selected from hydrogen, OH , C_{1-18} alkyl, C_{2-18} alkenyl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, C_{1-18} alkoxy, $-\text{NR}^{15}\text{R}^{16}$, aryl, an amino acid residue linked through an amino group of the amino acid, $\text{CH}_2\text{OCH}(=\text{O})\text{R}^{9a}$, or $\text{CH}_2\text{OC}(=\text{O})\text{OR}^{9a}$ where R^{9a} is C_{1-12} alkyl, C_{6-20} aryl, C_{6-20} alkylaryl or C_{6-20} aralkyl;

20 R^{10} and R^{11} are independently selected from the group consisting of hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, aryl, $-\text{C}(=\text{O})\text{R}^{12}$, heterocyclic, or an amino acid residue;

25 R^{12} is selected from the group consisting of hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, aryl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, or an amino acid residue;

30 R^{15} and R^{16} are independently selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, aryl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, or an amino acid residue;

35 R^{17} is independently selected from the group consisting of hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, C_{1-18} alkylsulfoxide, C_{1-18} alkylsulfone, C_{1-18} halogenated alkyl, C_{2-18} halogenated alkenyl, C_{2-18} halogenated alkynyl, C_{1-18} halogenated alkoxy, C_{1-18} halogenated alkylthio, C_{3-10} cycloalkyl, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, halogen, OH , CN , CO_2H , CO_2R^{18} , NO_2 , NR^7R^8 , haloalkyl, $\text{C}(=\text{O})\text{R}^{18}$, $\text{C}(=\text{S})\text{R}^{18}$, SH , aryl, aryloxy, arylthio, arylsulfoxide, arylsulfone, arylsulfonamide, arylalkyl, arylalkyloxy, arylalkylthio, heterocyclic, C_{1-18} hydroxyalkyl, where each of said aryl, aryloxy, arylthio, arylsulfoxide, arylsulfone, arylsulfonamide, arylalkyl, arylalkyloxy, arylalkylthio, heterocycle, or C_{1-18} hydroxyalkyl is optionally substituted with 1 or more R^{19} ;

5 R¹⁹ is selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, C₁₋₁₈ alkoxy, C₂₋₁₈ alkenyloxy, C₂₋₁₈ alkynyloxy, C₁₋₁₈ alkylthio, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, C₄₋₁₀ cycloalkynyl, halogen, -OH, -CN, cyanoalkyl, -NO₂, -NR²⁰R²¹, C₁₋₁₈ haloalkyl, C₁₋₁₈ haloalkyloxy, -C(=O)R¹⁸, -C(=O)OR¹⁸, -OalkenylC(=O)OR¹⁸, -OalkylC(=O)NR²⁰R²¹, -OalkylOC(=O)R¹⁸, -C(=S)R¹⁸, SH, -C(=O)N(C₁₋₆ alkyl), 10 -N(H)S(O)(O)(C₁₋₆ alkyl), aryl, heterocyclic, C₁₋₁₈alkylsulfone, arylsulfoxide, arylsulfonamide, aryl(C₁₋₁₈)alkyloxy, aryloxy, aryl(C₁₋₁₈ alkyl)oxy, arylthio, aryl(C₁₋₁₈)alkylthio or aryl(C₁₋₁₈)alkyl, where each may be optionally substituted with 1 or more =O, NR²⁰R²¹, CN, C₁₋₁₈ alkoxy, heterocyclic, C₁₋₁₈ haloalkyl, heterocyclic alkyl, heterocyclic connected to R¹⁷ by alkyl, alkoxyalkoxy or halogen;

15 R²⁰ and R²¹ are independently selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, -C(=O)R¹², carboxylester-substituted heterocyclic or -C(=S)R¹²;

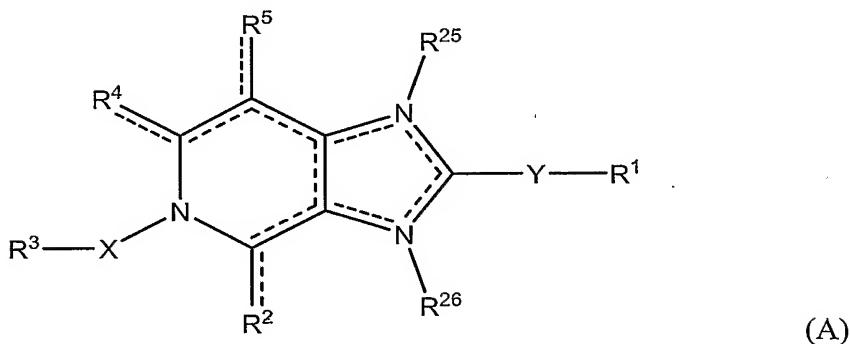
20 R²⁵ and R²⁶ are not present, or are independently selected from hydrogen, C₁₋₁₈ alkyl, C₃₋₁₀ cycloalkyl, aryl, heterocyclic, where each is optionally independently substituted with 1 to 4 of C₁₋₆ alkyl, C₁₋₆ alkoxy, halo, CH₂OH, benzyloxy, and OH; and

25 R²⁷ is selected from hydrogen, C₁₋₁₈ alkyl, C₃₋₁₀ cycloalkyl, (C₃₋₁₀ cycloalkyl)-C₁₋₆ alkyl, aryl, and aryl C₁₋₁₈ alkyl, and

the salts, tautomers, isomers and solvates thereof.

25

3. A compound having the general formula (A),



wherein:

5 the dotted lines represent an optional double bond, provided that no two double bonds are adjacent to one another, and that the dotted lines represent at least 3, optionally 4 double bonds;

10 R^1 is selected from hydrogen, aryl, heterocyclic, C_{1-10} alkoxy, C_{1-10} thioalkyl, $C_{1-C_{10}}$ alkyl-amino, $C_{1-C_{10}}$ dialkyl-amino, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, and C_{4-10} cycloalkynyl, wherein each are optionally substituted with 1 or more R^6 ;

15 Y is selected from single bond, O, $S(O)_m$, NR^{11} , or C_{1-10} alkylene, C_{2-10} alkenylene, C_{2-10} alkynylene, wherein each may optionally include 1 to 3 heteroatoms selected from O, S or N;

20 R^2 and R^4 are independently selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, halogen, -OH, -CN, -NO₂, -NR⁷R⁸, haloalkyloxy, haloalkyl, -C(=O)R⁹, -C(=S)R⁹, SH, aryl, aryloxy, arylthio, arylalkyl, C_{1-18} hydroxyalkyl, C_{3-10} cycloalkyl, C_{3-10} cycloalkyloxy, C_{3-10} cycloalkylthio, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, or heterocyclic, provided that when one of R^{25} or R^{26} is present, then either R^2 or R^4 is selected from (=O), (=S), and =NR²⁷;

25 X is selected from $C_{1-C_{10}}$ alkylene, C_{2-10} alkenylene or C_{2-10} alkynylene, where each may include one or more heteroatoms selected from O, S, or N, provided any such heteroatom is not adjacent to the N in the ring;

m is any integer from 0 to 2;

30 R^3 is a heterocycle optionally substituted with at least one R^{17} , provided R^3 -M-Q is not biphenyl;

R^5 is selected from hydrogen; C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, halogen, -OH, -CN, -NO₂, -NR⁷R⁸, haloalkyloxy, haloalkyl, -C(=O)R⁹, -C(=O)OR⁹, -C(=S)R⁹, SH, aryl, aryloxy, arylthio, arylalkyl, C_{1-18} hydroxyalkyl, C_{3-10} cycloalkyl, C_{3-10} cycloalkyloxy, C_{3-10} cycloalkylthio, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, or heterocyclic;

35 R^6 is selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, C_{1-18} alkylsulfoxide, C_{1-18} alkylsulfone, C_{1-18} halo-alkyl, C_{2-18} halo-alkenyl, C_{2-18} halo-alkynyl, C_{1-18} halo-alkoxy, C_{1-18} halo-alkylthio, C_{3-10} cycloalkyl, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, halogen, OH, CN, cyanoalkyl, -CO₂R¹⁸, NO₂, -NR⁷R⁸, C_{1-18} haloalkyl, C(=O)R¹⁸, C(=S)R¹⁸, SH, aryl, aryloxy,

5 arylthio, arylsulfoxide, arylsulfone, arylsulfonamide, aryl(C₁₋₁₈)alkyl, aryl(C₁₋₁₈)alkyloxy, aryl(C₁₋₁₈)alkylthio, heterocyclic, C₁₋₁₈ hydroxyalkyl, where each may be optionally substituted with at least 1 R¹⁹;

10 R⁷ and R⁸ are independently selected from hydrogen, C₁₋₁₈ alkyl, C₁₋₁₈ alkenyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, heterocyclic, -C(=O)R¹²; -C(=S)R¹², an amino acid residue linked through a carboxyl group thereof, or where R⁷ and R⁸ together with the nitrogen form a heterocyclic;

15 R⁹ and R¹⁸ are independently selected from hydrogen, OH, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, C₁₋₁₈ alkoxy, -NR¹⁵R¹⁶, aryl, an amino acid residue linked through an amino group of the amino acid, CH₂OCH(=O)R^{9a}, or CH₂OC(=O)OR^{9a} where R^{9a} is C₁-C₁₂ alkyl, C₆-C₂₀ aryl, C₆-C₂₀ alkylaryl or C₆-C₂₀ aralkyl;

20 R¹⁰ and R¹¹ are independently selected from the group consisting of hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, aryl, -C(=O)R¹², heterocyclic, or an amino acid residue;

25 R¹² is selected from the group consisting of hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, or an amino acid residue;

R¹⁵ and R¹⁶ are independently selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, or an amino acid residue;

30 R¹⁷ is M-Q-, wherein M is a C₃₋₁₀ cycloalkyl optionally substituted with 1 or more R¹⁹, and Q is a bond, or C₁₋₁₀ alkyl optionally substituted with 1 or more R¹⁹;

R¹⁹ is selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, C₁₋₁₈ alkoxy, C₂₋₁₈ alkenyloxy, C₂₋₁₈ alkynyloxy, C₁₋₁₈ alkylthio, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, C₄₋₁₀ cycloalkynyl, halogen, -OH, -CN, cyanoalkyl, -NO₂, -NR²⁰R²¹,

35 C₁₋₁₈ haloalkyl, C₁₋₁₈ haloalkyloxy, -C(=O)R¹⁸, -C(=O)OR¹⁸, -OalkenylC(=O)OR¹⁸, -OalkylC(=O)NR²⁰R²¹, -OalkylOC(=O)R¹⁸, -C(=S)R¹⁸, SH, -C(=O)N(C₁₋₆ alkyl), -N(H)S(O)(O)(C₁₋₆ alkyl), aryl, heterocyclic, C₁₋₁₈alkylsulfone, arylsulfoxide, arylsulfonamide, aryl(C₁₋₁₈)alkyloxy, aryloxy, aryl(C₁₋₁₈ alkyl)oxy, arylthio, aryl(C₁₋₁₈)alkylthio or aryl(C₁₋₁₈)alkyl, where each may be optionally substituted with 1 or more =O, NR²⁰R²¹, CN, C₁₋₁₈ alkoxy, heterocyclic, C₁₋₁₈ haloalkyl, heterocyclic alkyl, heterocyclic connected to R¹⁷ by alkyl, alkoxyalkoxy or halogen;

5 R²⁰ and R²¹ are independently selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, -C(=O)R¹², or -C(=S)R¹²;

10 R²⁵ and R²⁶ are not present, or are independently selected from hydrogen, C₁₋₁₈ alkyl, C₃₋₁₀ cycloalkyl, aryl, heterocyclic, where each is optionally independently substituted with 1 to 4 of C₁₋₆ alkyl, C₁₋₆ alkoxy, halo, CH₂OH, benzyloxy, and OH; and

15 R²⁷ is selected from hydrogen, C₁₋₁₈ alkyl, C₃₋₁₀ cycloalkyl, (C₃₋₁₀ cycloalkyl)-C₁₋₆ alkyl, aryl, and aryl C₁₋₁₈ alkyl, and
the salts, tautomers, isomers and solvates thereof.

15 4. The compound of claim 1, 2 or 3 wherein R³ is heterocycle.

20 5. The compound of claims 1, 2 or 3 wherein YR¹ is halophenyl.

25 6. The compound of claim 5 wherein halophenyl is ortho-fluorophenyl.

7. The compound of claims 1, 2 or 3 wherein R³ is isoxazolyl substituted with 1 R¹⁷.

25 8. The compound of claims 1, 2 or 3 wherein R¹⁷ is aryl or an aromatic heterocycle which is substituted with 1, 2 or 3 R¹⁹.

30 9. The compound of claims 1, 2 or 3 wherein YR¹ is none of hydrogen, an unsubstituted C₃₋₁₀ cycloalkyl, or C₁₋₆ alkyl.

30 10. The compound of claim 9 wherein YR¹ is not hydrogen.

35 11. The compound of claims 1, 2 or 3 wherein R¹⁹ is trihalomethyl, trihalomethoxy, alkoxy or halogen.

5 12. The compound of claims 1, 2 or 3 wherein R¹ is aryl or aromatic heterocycle substituted with 1, 2 or 3 R⁶ wherein R⁶ is halogen, C₁₋₁₈ alkoxy; or C₁₋₁₈ haloalkyl.

13. The compound of claims 12 wherein R¹ is phenyl substituted with 1, 2 or 3 halogens.

10

14. The compound of claims 1, 2 or 3 wherein halogen is fluoro.

15. The compound of claims 1, 2 or 3 wherein Y is a single bond, O, C₁₋₆ alkylene, C₂₋₆ alkenylene, C₂₋₆ alkynylene or one of said groups containing 1 to 3 heteroatoms selected from O, S or NR¹¹.

16. The compound of claim 15 wherein Y is -O(CH₂)₁₋₅-, -(CH₂)₁₋₄-O-(CH₂)₁₋₄-, -S-(CH₂)₁₋₅-, -(CH₂)₁₋₄-S-(CH₂)₁₋₄-, -NR¹¹-(CH₂)₁₋₅-, -(CH₂)₁₋₄-NR¹¹-(CH₂)₁₋₄ or C₃₋₁₀ cycloalkylidene.

20

17. The compound of claim 15 wherein Y is -OCH₂-, -CH₂O-, C₁₋₂ alkylene, C₂₋₃ alkenylene, C₂₋₃ alkynylene, O or a bond.

18. The compound of claim 15 wherein Y is a bond.

25

19. The compound of claims 1, 2 or 3 wherein YR¹ is not any one of H, an unsubstituted C₃₋₁₀ cycloalkyl or C1-C6 alkyl.

20. The compound of claims 1, 2 or 3 wherein YR¹ is not H.

30

21. The compound of claims 1, 2 or 3 wherein YR¹ is halo or halomethyl-substituted phenyl.

35 22. The compound of claims 1, 2 or 3 wherein halo or halomethyl are ortho or

meta.

5 23. The compound of claims 1, 2 or 3 wherein X is selected from the group consisting of alkylene, alkynylene or alkenylene and said hydrocarbons having an intrachain N, O or S heteroatom.

24. The compound of claims 1, 2 or 3 wherein X is alkyl.

10 25. The compound of claim 23 wherein X is selected from the group consisting of -CH₂-, -CH(CH₃)-, -CH₂-CH₂-, -CH₂-CH₂-CH₂-, -CH₂-CH₂-CH₂-CH₂-, -(CH₂)₂₋₄-O-(CH₂)₂₋₄-, -(CH₂)₂₋₄-S-(CH₂)₂₋₄-, -(CH₂)₂₋₄-NR¹⁰-(CH₂)₂₋₄-, C₃₋₁₀ cycloalkylidene, C₂₋₆ alkenylene and C₂₋₆ alkynylene.

15 26. The compound of claims 1, 2 or 3 wherein X is methylene.

27. The compound of claims 1, 2 or 3 wherein R³ is aryl or a heterocycle substituted with 0 to 3 R¹⁷.

20 28. The compound of claim 27 wherein the heterocycle is an aromatic heterocycle.

25 29. The compound of claim 28 wherein the heterocycle contains 1, 2 or 3 N, S or O atoms in the ring, is linked to X through a ring carbon atom and contains 4 to 6 total ring atoms.

30 30. The compound of claims 1, 2 or 3 wherein R³ is isoxazolyl substituted with 1 to 3 R¹⁷.

30 31. The compound of claims 1, 2 or 3 wherein R¹⁷ is aryl or a heterocycle further substituted with 1 to 3 R¹⁹.

35 32. The compound of claims 1 or 3 wherein M is aryl or aromatic heterocycle.

5 33. The compound of claims 1 or 3 wherein Q contains 0 to 20 atoms selected
from C, O, S, N and H.

34. The compound of claims 1 or 3 wherein M is a cyclic group selected from R¹⁷.

10 35. The compound of claim 2 wherein R¹⁷ is selected from the group consisting of
C₃₋₁₀ cycloalkyl, C₃₋₁₀ cycloalkenyl, C₇₋₁₀ cycloalkynyl, halogen, aryl, aryloxy,
arylthio, arylsulfoxide, arylsulfone, arylsulfonamide, arylalkyl; arylalkyloxy;
arylalkylthio; heterocycle; C₁₋₁₈ hydroxyalkyl, each of said C₃₋₁₀ cycloalkyl, C₃₋₁₀
cycloalkenyl, C₇₋₁₀ cycloalkynyl, halogen, aryl, aryloxy, arylthio, arylsulfoxide,
15 arylsulfone, arylsulfonamide, arylalkyl; arylalkyloxy; arylalkylthio; heterocycle; and
C₁₋₁₈ hydroxyalkyl is unsubstituted or is substituted 1 or more R¹⁹.

20 36. The compound of claim 2 wherein R¹⁷ is selected from the group consisting
of aryl and heterocycle, and where said aryl or heterocycle is optionally substituted
with 1 or more R¹⁹.

37. The compound of claims 1, 2 or 3 wherein R⁹ and R¹⁸ are H, OH or alkyl.

25 38. The compound of claims 1, 2 or 3 wherein R⁵ is H.

39. The compound of claims 1, 2 or 3 wherein R⁶ is halogen.

40. The compound of claims 1, 2 or 3 wherein R⁷, R⁸, R¹⁰, R¹¹, R¹⁵, R¹⁶, R²⁰, and
R²¹ are independently H or C₁₋₁₈ alkyl.

30 41. The compound of claims 1, 2 or 3 wherein R¹² is OH or alkyl.

42. The compound of claims 1, 2 or 3 wherein R¹⁹ is selected from the group
consisting of H; C₁₋₁₈ alkyl; C₂₋₁₈ alkenyl; C₂₋₁₈ alkynyl; C₁₋₁₈ alkoxy; alkenyloxy;
35 alkynyloxy; C₁₋₁₈ alkylthio; C₃₋₁₀ cycloalkyl; C₄₋₁₀ cycloalkenyl; C₄₋₁₀ cycloalkynyl;
halogen; OH; CN; cyanoalkyl; NO₂; NR²⁰R²¹; haloalkyl; haloalkyloxy; C(=O)R¹⁸;

5 C(=O)OR¹⁸; OalkenylC(=O)OR¹⁸; -OalkylC(=O)NR²⁰R²¹; aryl; heterocycle; -
 OalkylOC(=O)R¹⁸; C(=O)N(C₁₋₆ alkyl), N(H)S(O)(O)(C₁₋₆ alkyl); arylalkyloxy;
 aryloxy; arylalkyloxy; and arylalkyl; each of which is unsubstituted or substituted
 with 1 or more =O; NR²⁰R²¹; CN; alkoxy; heterocycle; haloalkyl- or alkyl-
 substituted heterocycle; and heterocycle linked to R¹⁷ by alkyl; alkoxyalkoxy or
 10 halogen.

43. The compound of claim 42 wherein R¹⁹ is independently selected from the
 group consisting of halogen, N(R²⁰ R²¹), alkoxy. halo-substituted alkyl and halo-
 substituted alkoxy.

15

44. The compound of claims 1, 2 or 3 wherein R²⁵ and R²⁶ are not present.

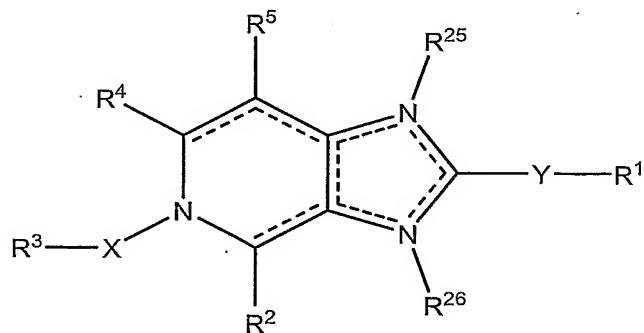
45. The compound of claims 1, 2 or 3 which is not substituted at R²⁵ but is
 substituted at R²⁶, and either R² or R⁴ is selected from (=O), (=S), and (=NR²⁷).

20

46. The compound of claims 1, 2 or 3 wherein haloalkyl or haloalkyloxy is -CF₃
 or -OCF₃.

47. A composition comprising a pharmaceutically acceptable excipient and a
 25 compound of claims 1, 2 or 3.

48. A compound having the general formula (B),



(B)

30 wherein:

5 the dotted lines represent an optional double bond, provided that no two double bonds are adjacent to one another, and that the dotted lines represent at least 3, optionally 4 double bonds;

10 R^1 is selected from hydrogen, aryl, heterocyclic, C_{1-10} alkoxy, C_{1-10} thioalkyl, C_{1-10} alkyl-amino, C_{1-10} dialkyl-amino, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, and C_{4-10} cycloalkynyl, wherein each are optionally substituted with 1 or more R^6 ;

15 Y is selected from single bond, O , $S(O)_m$, NR^{11} , or C_{1-10} alkylene, C_{2-10} alkenylene, C_{2-10} alkynylene, wherein each may optionally include 1 to 3 heteroatoms selected from O , S or N ;

20 R^2 and R^4 are independently selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, halogen, $-OH$, $-CN$, $-NO_2$, $-NR^7R^8$, haloalkyloxy, haloalkyl, $-C(=O)R^9$, $-C(=S)R^9$, SH , aryl, aryloxy, arylthio, arylalkyl, C_{1-18} hydroxyalkyl, C_{3-10} cycloalkyl, C_{3-10} cycloalkyloxy, C_{3-10} cycloalkylthio, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, or heterocyclic, provided that when one of R^{25} or R^{26} is present, then either R^2 or R^4 is selected from $(=O)$, $(=S)$, and $=NR^{27}$;

25 X is selected from C_{1-10} alkylene, C_{2-10} alkenylene or C_{2-10} alkynylene, where each may include one or more heteroatoms selected from O , S , or N , provided any such heteroatom is not adjacent to the N in the ring;

m is any integer from 0 to 2;

30 R^3 is selected from aryl, aryloxy, arylthio, cycloalkyl, cycloalkenyl, cycloalkynyl, aryl- $N(R^{10})$ -, or heterocyclic, where each said substituent may be optionally substituted with at least one R^{17} , provided that for cycloalkenyl the double bond is not adjacent to a nitrogen, and provided R^3 M-Q- is not biphenyl;

35 R^5 is selected from hydrogen; C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, halogen, $-OH$, $-CN$, $-NO_2$, $-NR^7R^8$, haloalkyloxy, haloalkyl, $-C(=O)R^9$, $-C(=O)OR^9$, $-C(=S)R^9$, SH , aryl, aryloxy, arylthio, arylalkyl, C_{1-18} hydroxyalkyl, C_{3-10} cycloalkyl, C_{3-10} cycloalkyloxy, C_{3-10} cycloalkylthio, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, or heterocyclic;

40 R^6 is selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, C_{1-18} alkylsulfoxide, C_{1-18} alkylsulfone, C_{1-18} halo-alkyl, C_{2-18} halo-alkenyl, C_{2-18} halo-alkynyl, C_{1-18} halo-alkoxy, C_{1-18} halo-alkylthio, C_{3-10}

5 cycloalkyl, C₃₋₁₀ cycloalkenyl, C₇₋₁₀ cycloalkynyl, halogen, OH, CN, cyanoalkyl, -CO₂R¹⁸, NO₂, -NR⁷R⁸, C₁₋₁₈ haloalkyl, C(=O)R¹⁸, C(=S)R¹⁸, SH, aryl, aryloxy, arylthio, arylsulfoxide, arylsulfone, arylsulfonamide, aryl(C₁₋₁₈)alkyl, aryl(C₁₋₁₈)alkyloxy, aryl(C₁₋₁₈)alkylthio, heterocyclic, C₁₋₁₈ hydroxyalkyl, where each may be optionally substituted with at least 1 R¹⁹;

10 R⁷ and R⁸ are independently selected from hydrogen, C₁₋₁₈ alkyl, C₁₋₁₈ alkenyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, heterocyclic, -C(=O)R¹²; -C(=S)R¹², an amino acid residue linked through a carboxyl group thereof, or where R⁷ and R⁸ together with the nitrogen form a heterocyclic;

15 R⁹ and R¹⁸ are independently selected from hydrogen, OH, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, C₁₋₁₈ alkoxy, -NR¹⁵R¹⁶, aryl, an amino acid residue linked through an amino group of the amino acid, CH₂OCH(=O)R^{9a}, or CH₂OC(=O)OR^{9a} where R^{9a} is C_{1-C₁₂} alkyl, C_{6-C₂₀} aryl, C_{6-C₂₀} alkylaryl or C_{6-C₂₀} aralkyl;

20 R¹⁰ and R¹¹ are independently selected from the group consisting of hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, aryl, -C(=O)R¹², heterocyclic, or an amino acid residue;

25 R¹² is selected from the group consisting of hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, or an amino acid residue;

30 R¹⁵ and R¹⁶ are independently selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, or an amino acid residue;

35 R¹⁷ is independently M-Q- wherein M is a ring optionally substituted with 1 or more R¹⁹, and Q is a bond or a linking group connecting M to R³ having 1 to 10 atoms and optionally substituted with 1 or more R¹⁹;

30 R¹⁹ is selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, C₁₋₁₈ alkoxy, C₂₋₁₈ alkenyloxy, C₂₋₁₈ alkynyloxy, C₁₋₁₈ alkylthio, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, C₄₋₁₀ cycloalkynyl, halogen, -OH, -CN, cyanoalkyl, -NO₂, -NR²⁰R²¹, C₁₋₁₈ haloalkyl, C₁₋₁₈ haloalkyloxy, -C(=O)R¹⁸, -C(=O)OR¹⁸, -OalkenylC(=O)OR¹⁸, -OalkylC(=O)NR²⁰R²¹, -OalkylOC(=O)R¹⁸, -C(=S)R¹⁸, SH, -C(=O)N(C₁₋₆ alkyl), -N(H)S(O)(O)(C₁₋₆ alkyl), aryl, heterocyclic, C₁₋₁₈ alkylsulfone, arylsulfoxide, arylsulfonamide, aryl(C₁₋₁₈)alkyloxy, aryloxy, aryl(C₁₋₁₈ alkyl)oxy, arylthio, aryl(C₁₋₁₈)alkylthio, heterocyclic, C₁₋₁₈ hydroxyalkyl, where each may be optionally substituted with at least 1 R¹⁹;

5 18) alkylthio or aryl(C₁₋₁₈)alkyl, where each may be optionally substituted with 1 or
more =O, NR²⁰R²¹, CN, C₁₋₁₈ alkoxy, heterocyclic, C₁₋₁₈ haloalkyl, heterocyclic alkyl,
heterocyclic connected to R¹⁷ by alkyl, alkoxyalkoxy or halogen;

10 R²⁰ and R²¹ are independently selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈
alkenyl, C₂₋₁₈ alkynyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, -C(=O)R¹², or
-C(=S)R¹²;

R²⁷ is selected from hydrogen, C₁₋₁₈ alkyl, C₃₋₁₀ cycloalkyl, (C₃₋₁₀ cycloalkyl)-
C₁₋₆ alkyl, aryl, and aryl C₁₋₁₈ alkyl, and

salts, tautomers, isomers and solvates thereof.

49. The compound of claim 48 wherein Y is a single bond, and R¹ is aryl.

15

50. The compound of claim 48 wherein X is C_{1-C10} alkylene, C₂₋₁₀ alkenylene or
C₂₋₁₀ alkynylene.

51. The compound of claim 48 wherein R³ is heterocyclic.

20

52. The compound of claim 48 wherein R³ is heterocyclic substituted with R¹⁷
where Q is a bond and M is aryl.

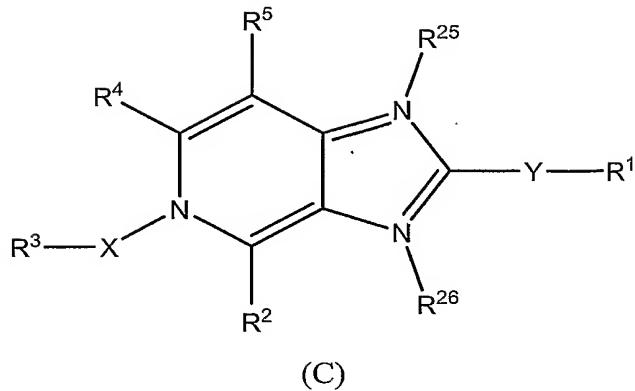
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54. The compound of claim 48 wherein R³ is isoxazole substituted with R¹⁷ where
Q is a bond and M is aryl.

30

55. The compound of claim 48 wherein R³ is isoxazole substituted with R¹⁷ where
Q is a bond and M is phenyl.

56. A compound having the general formula (C),



wherein:

the dotted lines represent an optional double bond, provided that no two double bonds are adjacent to one another, and that the dotted lines represent at least 3, 10 optionally 4 double bonds;

15 R^1 is selected from hydrogen, aryl, heterocyclic, C_{1-10} alkoxy, C_{1-10} thioalkyl, C_{1-10} alkyl-amino, C_{1-10} dialkyl-amino, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, and C_{4-10} cycloalkynyl, wherein each are optionally substituted with 1 or more R^6 ;

20 Y is selected from single bond, O , $S(O)_m$, NR^{11} , or C_{1-10} alkylene, C_{2-10} alkenylene, C_{2-10} alkynylene, wherein each may optionally include 1 to 3 heteroatoms selected from O , S or N ;

R^2 and R^4 are independently selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, halogen, $-OH$, $-CN$, $-NO_2$, $-NR^7R^8$, haloalkyloxy, haloalkyl, $-C(=O)R^9$, $-C(=S)R^9$, SH , aryl, aryloxy, arylthio, arylalkyl, C_{1-18} hydroxyalkyl, C_{3-10} cycloalkyl, C_{3-10} cycloalkyloxy, C_{3-10} cycloalkylthio, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, or heterocyclic, provided that when one of R^{25} or R^{26} is present, then either R^2 or R^4 is selected from $(=O)$, $(=S)$, and $=NR^{27}$;

25 X is selected from C_{1-10} alkylene, C_{2-10} alkenylene or C_{2-10} alkynylene, where each may include one or more heteroatoms selected from O , S , or N , provided any such heteroatom is not adjacent to the N in the ring;

m is any integer from 0 to 2;

30 R^3 is selected from aryl, aryloxy, arylthio, cycloalkyl, cycloalkenyl, cycloalkynyl, aryl- $N(R^{10})-$, or heterocyclic, where each said substituent may be optionally substituted with at least one R^{17} , provided that for cycloalkenyl the double bond is not adjacent to a nitrogen, and provided R^3 M-Q- is not biphenyl;

5 R⁵ is selected from hydrogen; C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, C₁₋₁₈ alkoxy, C₁₋₁₈ alkylthio, halogen, -OH, -CN, -NO₂, -NR⁷R⁸, haloalkyloxy, haloalkyl, -C(=O)R⁹, -C(=O)OR⁹, -C(=S)R⁹, SH, aryl, aryloxy, arylthio, arylalkyl, C₁₋₁₈ hydroxyalkyl, C₃₋₁₀ cycloalkyl, C₃₋₁₀ cycloalkyloxy, C₃₋₁₀ cycloalkylthio, C₃₋₁₀ cycloalkenyl, C₇₋₁₀ cycloalkynyl, or heterocyclic;

10 R⁶ is selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, C₁₋₁₈ alkoxy, C₁₋₁₈ alkylthio, C₁₋₁₈ alkylsulfoxide, C₁₋₁₈ alkylsulfone, C₁₋₁₈ halo-alkyl, C₂₋₁₈ halo-alkenyl, C₂₋₁₈ halo-alkynyl, C₁₋₁₈ halo-alkoxy, C₁₋₁₈ halo-alkylthio, C₃₋₁₀ cycloalkyl, C₃₋₁₀ cycloalkenyl, C₇₋₁₀ cycloalkynyl, halogen, OH, CN, cyanoalkyl, -CO₂R¹⁸, NO₂, -NR⁷R⁸, C₁₋₁₈ haloalkyl, C(=O)R¹⁸, C(=S)R¹⁸, SH, aryl, aryloxy, 15 arylthio, arylsulfoxide, arylsulfone, arylsulfonamide, aryl(C₁₋₁₈)alkyl, aryl(C₁₋₁₈)alkyloxy, aryl(C₁₋₁₈)alkylthio, heterocyclic, C₁₋₁₈ hydroxyalkyl, where each may be optionally substituted with at least 1 R¹⁹;

20 R⁷ and R⁸ are independently selected from hydrogen, C₁₋₁₈ alkyl, C₁₋₁₈ alkenyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, heterocyclic, -C(=O)R¹²; -C(=S)R¹², an amino acid residue linked through a carboxyl group thereof, or where R⁷ and R⁸ together with the nitrogen form a heterocyclic;

25 R⁹ and R¹⁸ are independently selected from hydrogen, OH, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, C₁₋₁₈ alkoxy, -NR¹⁵R¹⁶, aryl, an amino acid residue linked through an amino group of the amino acid, CH₂OCH(=O)R^{9a}, or CH₂OC(=O)OR^{9a} where R^{9a} is C_{1-C12} alkyl, C_{6-C20} aryl, C_{6-C20} alkylaryl or C_{6-C20} aralkyl;

30 R¹⁰ and R¹¹ are independently selected from the group consisting of hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, aryl, -C(=O)R¹², heterocyclic, or an amino acid residue;

 R¹² is selected from the group consisting of hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, or an amino acid residue;

 R¹⁵ and R¹⁶ are independently selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, or an amino acid residue;

5 R^{17} is independently M-Q- wherein M is a ring optionally substituted with 1 or more R^{19} , and Q is a bond or a linking group connecting M to R^3 having 1 to 10 atoms and optionally substituted with 1 or more R^{19} ;

10 R^{19} is selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{2-18} alkenyloxy, C_{2-18} alkynyloxy, C_{1-18} alkylthio, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, C_{4-10} cycloalkynyl, halogen, -OH, -CN, cyanoalkyl, -NO₂, -NR²⁰R²¹, C_{1-18} haloalkyl, C_{1-18} haloalkyloxy, -C(=O)R¹⁸, -C(=O)OR¹⁸, -OalkenylC(=O)OR¹⁸, -OalkylC(=O)NR²⁰R²¹, -OalkylOC(=O)R¹⁸, -C(=S)R¹⁸, SH, -C(=O)N(C_{1-6} alkyl), -N(H)S(O)(O)(C_{1-6} alkyl), aryl, heterocyclic, C_{1-18} alkylsulfone, arylsulfoxide, arylsulfonamide, aryl(C_{1-18})alkyloxy, aryloxy, aryl(C_{1-18} alkyl)oxy, arylthio, aryl(C_{1-18})alkylthio or aryl(C_{1-18})alkyl, where each may be optionally substituted with 1 or more =O, NR²⁰R²¹, CN, C_{1-18} alkoxy, heterocyclic, C_{1-18} haloalkyl, heterocyclic alkyl, heterocyclic connected to R^{17} by alkyl, alkoxyalkoxy or halogen;

15 R^{20} and R^{21} are independently selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, aryl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, -C(=O)R¹², or -C(=S)R¹²;

20 R^{27} is selected from hydrogen, C_{1-18} alkyl, C_{3-10} cycloalkyl, (C_{3-10} cycloalkyl)- C_{1-6} alkyl, aryl, and aryl C_{1-18} alkyl, and salts, tautomers, isomers and solvates thereof.

25 57. The compound of claim 56 wherein Y is a single bond, and R^1 is aryl.

30 58. The compound of claim 56 wherein X is $C_{1-C_{10}}$ alkylene, C_{2-10} alkenylene or C_{2-10} alkynylene.

35 59. The compound of claim 56 wherein R^3 is heterocyclic.

60. The compound of claim 56 wherein R^3 is heterocyclic substituted with R^{17} where Q is a bond and M is aryl.

35 61. The compound of claim 56 wherein Y is a single bond, and R^1 is phenyl.

5 62. The compound of claim 56 wherein R³ is isoxazole substituted with R¹⁷ where
Q is a bond and M is aryl.

63. The compound of claim 56 wherein R³ is isoxazole substituted with R¹⁷ where
Q is a bond and M is phenyl.

10 64. A method comprising administering to a subject in need of treatment or
prophylaxis of a viral infection an antivirally effective amount of a compound of
claims 1, 2, 3, 48 or 56.

15 65. The method of claim 64, wherein the viral infection is an infection of a
hepatitis-C virus.

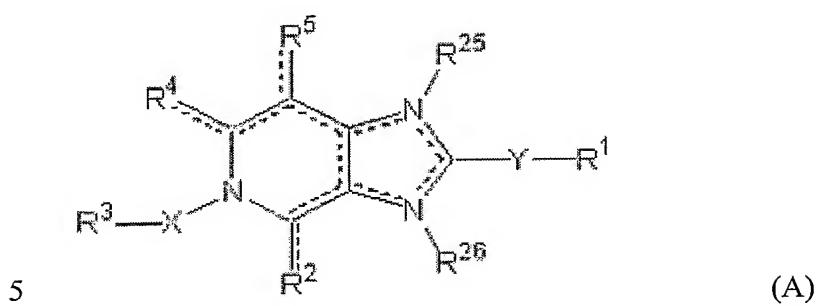
66. The method of claim 65 further comprising administering at least one
additional antiviral therapy to the subject.

20 67. The method of claim 66 wherein the additional therapy is selected from the
group consisting of an interferon alpha and ribavirin.

25 68. A method of screening antiviral compounds which comprises providing a
compound of claims 1, 2, 3, 48 or 56 and determining the anti-viral activity of said
compound.

69. The method of claim 68 wherein said anti-viral activity is determined by the
activity of said compound against one or more viruses belonging to the family of the
30 Flaviviridae and/or of the Picornaviridae.

70. A method for assaying the structure-activity of analogues of formula (A)
compounds



wherein the substituents are defined in WO 2004/005286, comprising

(c) preparing a compound of formula (A) in which at least one substituent is not disclosed by WO 2004/005286; and
10 (d) determining the anti-HCV activity of the compound of step (a).

71. The method of claim 70 wherein the substituent is located at R³, R², R⁴, R²⁶ and/or R⁵.

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